

Dietary Antioxidants and Asthma Risk

Antioxidant constituents of diet decrease **risk of asthma**.

General Information

Broad Focus Area	Asthma
Background and Justification	<p>Interest in the relation between dietary antioxidant consumption and risk of asthma arises from several directions. First, general population-based health surveys have demonstrated modest direct associations between pulmonary function and antioxidant consumption and serum levels.^{1,2} Second, markers of oxidative damage are found at higher concentrations in the exhaled breath condensate of children with asthma compared to those without asthma, while markers of antioxidant status are higher among children without asthma.³ Third, the deleterious pulmonary response to ozone exposure is somewhat modified by oral administration of Vitamins C and E.^{4,5} Pulmonary inflammatory response to ozone exposure is thought to be mediated at least in part by oxidative damage to airway tissues.</p> <p>Little direct evidence exists, however, concerning the relationship between antioxidant consumption or serum levels and the development of asthma, especially in relation to exposure to potential oxidative stressors such as ozone, nitrous oxides, or environmental tobacco smoke. In addition, examination of the temporal relationship between antioxidant consumption on subsequent outcome and assessment of genetic variation in immunologic response to oxidative stress and the potential modifying influence of antioxidant exposure will be important in understanding potential intervention techniques, whether related to asthma incidence or treatment.⁶</p>
Prevalence/ Incidence	<p>Nine million children less than 18 years of age are estimated to have asthma.⁷ Among children, it is the most common chronic illness.⁸ The prevalence of asthma increased from 35 to 62 per 1,000 children aged 0 to 17 years between 1980 and 1996.⁹ Exposure to bioaerosols and outdoor air pollution is ubiquitous but varies by location and time.</p>
Economic Impact	<p>In 1997, the annual estimated cost of pediatric asthma in the US was \$6.6 billion.¹⁰ By 2002, the total cost of asthma was estimated at \$14 billion.¹¹ The more severe forms of asthma account for a disproportionate amount of the total direct costs; one study estimated that less than 20% of asthmatics account for over 80% of the direct costs.¹² Asthma also poses a substantial and increasing public health burden in lost time from school and usual activities and in health care utilization.</p>

Exposure Measures		Outcome Measures	
<i>Primary/Maternal</i>	Antioxidant consumption and antioxidant levels	<i>Primary/Maternal</i>	
Methods	Interview/questionnaire (diet/nutrition assessment, vitamin use); blood samples; urine samples; breast milk	Methods	
Life Stage	Prenatal through infancy	Life Stage	
<i>Primary/Child</i>	Antioxidant consumption	<i>Primary/Child</i>	Decreased risk of asthma

	(maternal and child) and antioxidant levels			measured via allergy; asthma in index child; airway reactivity
Methods	Urine samples; blood samples; interview/questionnaire (diet/nutrition assessment, vitamin use)		Methods	Direct observation by medical professional; medical record review; interview/questionnaire; blood samples; urine samples
Life Stage	Periodic, 0-3 months to year 20		Life Stage	Periodic, birth to year 20

Important Confounders/Covariates	
Vitamins	Vitamin C at levels of 60/mg/day for nonsmokers and 100/mg/day for smokers may have a protective effect; there are still several unresolved questions. ^{13,14}
Smoking	Smoking, which is related to higher levels of oxidative stress, may reduce the effect of an antioxidative diet ^{13,15}
Red wine, apples	Red wine and apple consumption (in adults) was negatively associated with asthma severity (OR = 0.89) This may suggest a protective effect of flavonoids. ¹⁶

Population of Interest	Estimated Effect that is Detectable
All children	The smallest detectable relative risk is approximately 1.2. This power estimate assumes a sample size of 100,000 at age of diagnosis, an asthma incidence of 5%, and a cut-off value for “high” exposure based on the upper 5 th percentile of NCS subjects (i.e., a proportion exposed of 0.05). It assumes only a main effects model based on exposure to a single factor (e.g., a single pollutant) without consideration of interactions with other exposures, genetics, family history, etc. ¹⁷

Other Design Issues	
Ethical/Burden Considerations	Blood studies, especially fasting, in younger children will require careful attention.

References:

¹ Schunemann, H.J. et al. 2002. Lung function in relation to intake of carotenoids and other antioxidant vitamins in a population-based study. American Journal of Epidemiology 155: 463-471.

² Schwartz, J. and S.T. Weiss. 1990. Dietary factors and their relation to respiratory symptoms, The Second National Health and Nutrition Examination Survey. American Journal of Epidemiology 132: 67-76.

³ Corradi, M. et al. 2003. Aldehydes and glutathione in exhaled breath condensate of children with asthma exacerbation. American Journal of Respiratory and Critical Care Medicine 167(3): 395-399. Originally published online Oct 31 2002.

⁴ Samet, J.M. et al. 2001. Effect of antioxidant supplementation on ozone-induced lung injury in human subjects. American Journal of Respiratory and Critical Care Medicine 164: 819-825.

⁵ Romieu, I. et al. 1998. Antioxidant supplementation and respiratory functions among workers exposed to high levels of ozone. American Journal of Respiratory and Critical Care Medicine 158: 226-232.

- ⁶ NCS Interagency Coordinating Committee (ICC). Supporting documentation for the working list of NCS Core Hypotheses presented at the December, 2002 NCS Study Assembly meeting – Draft: “Rationale Document.” 14 February 2003.
- ⁷ Dey, A.N., Schiller, J.S., Tai, D.A. 2004. Summary Health Statistics for U.S. Children: National Health Interview Survey, 2002. Vital Health Stat 10 (221). National Center for Health Statistics, Centers for Disease Control and Prevention.
- ⁸ NAS. 2000. Clearing the Air: Asthma and Indoor Air Exposures. National Academy of Sciences Institute of Medicine, Division of Health Promotion and Disease Prevention. National Academy Press, Washington, D.C. 438 pp.
- ⁹ NCHS. 1979 through 1999. “Current Estimates from the National Health Interview Survey.” Vital and Health Statistics Series 10.
- ¹⁰ Landrigan, P.J., Schechter, C.B., Lipton, J.M., Fahs, M.C., Schwartz, J. 2002. Environmental pollutants and disease in American children: estimates of morbidity, mortality, and costs for lead poisoning, asthma, cancer, and developmental disabilities. Environmental Health Perspectives 110(7): 721-728.
- ¹¹ American Lung Association. March 2003. Trends in Asthma Morbidity and Mortality. Epidemiology & Statistics Unit.
- ¹² Weiss, K.B. 2001. The health economics of asthma and rhinitis. I. Assessing the economic impact. Journal of Allergy & Clinical Immunology 107(1): 3-8.
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- ¹⁴ Smit, H.A., Grievink, L., et al. 1999. Proc Nutr Soc. 58(2): 309-19.
- ¹⁵ Farchi, S., Forastiere, F., et al. 2003. Dietary factors associated with wheezing and allergic rhinitis in children. Eur Respir J. 22(5): 772-80.
- ¹⁶ Shaheen, S.O., Sterne, J.A., et al. 2001. Dietary antioxidants and asthma in adults: population-based case-control study. Am J Respir Crit Care Med. 164(10 Pt 1): 1823-8.
- ¹⁷ NCS Interagency Coordinating Committee (ICC), 2003 (see above).